Title: NON-TOXIC MUTANTS OF PATHOGENIC GRAM-NEGATIVE BACTERIA

Dkt: 875.001US2

IN THE CLAIMS

The listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

Claims 1-21 (Cancelled)

- 12
- Claim 22. (Currently amended) A method of making a mutant endotoxin comprising mutating an htrB gene encoding a wild type endotoxin in a wild type gram-negative bacterial pathogen, wherein the wild type pathogen produces wild type endotoxin, wherein mutating the htrB gene provides to provide a mutant gram-negative bacterial pathogen that produces the mutant endotoxin; wherein the lipid A of the mutant endotoxin is the same as the lipid A of the wild type endotoxin except the lipid A of the mutant endotoxin for lacking lacks one or more secondary acyl chains as compared to the of lipid A of the wild type endotoxin, and wherein the mutant endotoxin has substantially reduced toxicity when compared to the wild type endotoxin of the wild type gram negative bacterial pathogen; and purifying the mutant endotoxin from the mutant gram-negative bacterial pathogen.
- Claim 23. (Currently amended) A The mutant endotoxin made according to the method of claim 22, wherein the mutant endotoxin was purified from the htrB gram-negative bacterial mutant pathogen by phenol-water extraction or by protease digestion.
- Claim 24. (Previously presented) The mutant endotoxin according to claim 23, wherein the mutant endotoxin is conjugated to a carrier protein.
- Claim 25. (Currently amended) A <u>The</u> mutant endotoxin made according to the method of claim 22.

Claim 26. (Previously presented) The mutant endotoxin according to claim 25, wherein the mutant endotoxin is conjugated to a carrier protein.

Claims 27-28 (Cancelled)

Claim 29. (Currently amended) A method for producing endotoxin-specific antisera, the method comprising

- immunizing an individual with a vaccine formulation comprising an htrB (a) mutant of a gram-negative bacterial pathogen, endotoxin isolated from the htrB mutant of the gram-negative bacterial pathogen, or endotoxin purified from the htrB mutant of the gram-negative bacterial pathogen wherein the endotoxin is conjugated to a carrier protein; and
- collecting antibody specific for the htrB mutant produced from the (b) immunized individual; wherein the lipid A of the htrB mutant endotoxin is the same as the lipid A of the wild type endotoxin except that the lipid A of the htrB mutant endotoxin lacks for lacking one or more secondary acyl chains as compared to the of lipid A of the

Claims 30-31 (Cancelled)

wild type endotoxin.

- Claim 32. (Previously presented) The method of claim 22 wherein the gram-negative bacterial pathogen is of the genera Haemophilus, Neisseria, Moraxella, Campylobacter, Shigella or Pseudomonas.
- Claim 33. (Previously presented) The method of claim 29 wherein the gram-negative bacterial pathogen is of the genera Haemophilus, Neisseria, Moraxella, Campylobacter, Shigella or Pseudomonas.

AMENDMENT AND RESPONSE UNDER 37 CFR 1.111

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Claim 34.

(Cancelled)

II Conclude